

WHAT IS CLAIMED IS:

1. An isolated variant allele of a human orphanin FQ/nociceptin receptor gene, comprising a DNA sequence having at least one variation in SEQ ID NO:1, wherein said variation comprises G-46A, GIVS I 135C, GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T, C1126G, or any combination thereof.
2. The isolated variant allele of Claim 1, detectably labeled.
3. The isolated variant allele of Claim 2, wherein said detectable label comprises a radioactive element, a chemical which fluoresces, or an enzyme.
4. An isolated nucleic acid molecule selectively hybridizable to the isolated variant allele of Claim 1.
5. The isolated nucleic acid molecule of Claim 4, detectably labeled.
6. The isolated nucleic acid molecule of Claim 5, wherein said detectable label comprises a radioactive element, a chemical that fluoresces, or an enzyme.
7. A cloning vector comprising an isolated variant allele of a human orphanin FQ/nociceptin receptor gene and an origin of replication, wherein said variant allele comprises a DNA sequence having at least one variation in SEQ ID NO:1, wherein said variation comprises G-46A, GIVS I 135C, GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T, C1126G, or any combination thereof.
8. A cloning vector comprising an origin of replication and an isolated nucleic acid molecule selectively hybridizable to an isolated variant allele of a human orphanin FQ/nociceptin receptor gene, wherein said variant allele comprises a DNA sequence having at least one variation in SEQ ID NO:1, wherein said at least one variation comprises G-46A, GIVS I 135C, GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T, C1126G, or any combination thereof.



comprises G-46A, GIVS I 135C, GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T, C1126G, or any combination thereof.

15. A unicellular host transformed with an expression vector comprising an isolated nucleic acid molecule selectively hybridizable to an isolated variant allele of a human orphanin FQ/nociceptin receptor gene, wherein said isolated nucleic acid molecule is operatively associated with a promoter, and said variant allele comprises a DNA sequence having at least one variation in SEQ ID NO:1, wherein said at least one variation comprises G-46A, GIVS I 135C, GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T, C1126G, or any combination thereof.
16. The unicellular host of either of Claims 14 or 15, wherein said host comprises *E. coli*, *Pseudomonas*, *Bacillus*, *Streptomyces*, yeast, CHO, R1.1, B-W, L-M, COS1, COS7, BSC1, BSC40, BMT10 or Sf9 cells.
17. A method for determining a susceptibility in a subject to at least one physiological response, condition or disease related to the endogenous opioid system, nociception, neurotransmitter release (including dopamine, GABA, noradrenaline, and serotonin), anxiety and stress, learning, memory and cognition, alcohol self-administration, behavioral sensitization to cocaine, drug addiction, opiate withdrawal and tolerance, food intake, immune function, cardiovascular function, renal function, gastrointestinal function, or motor function, comprising the steps of:
  - a) removing a bodily sample from said subject, wherein said sample comprises a first and second allele comprising a human orphanin FQ/nociceptin receptor gene;
  - b) determining whether said human orphanin FQ/nociceptin receptor gene of said first allele comprises a DNA sequence having at least one variation in SEQ ID NO:1, wherein said variation comprises G-46A, GIVS I 135C, GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T, C1126G, or any combination thereof, such that the presence of said at least one variation in said human orphanin FQ/nociceptin receptor gene of said first allele is expected to be indicative of the subject's susceptibility to at least one said physiological response, condition or disease relative to the susceptibility to said at least one

19 said physiological response, condition or disease in a standard.

- 1 18. The method for determining a susceptibility to at least one addictive disease of Claim 17,  
2 further comprising the step of determining whether said human orphanin FQ/nociceptin  
3 receptor gene of said second allele comprises a DNA sequence having at least one  
4 variation in SEQ ID NO:1, wherein said variation comprises G-46A, GIVS I 135C,  
5 GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T, C1126G, or any  
6 combination thereof, such that the presence of said at least one variation in said human  
7 orphanin FQ/nociceptin receptor gene of said second allele is expected to be indicative  
8 of the subject's susceptibility to said at least one physiological response, condition or  
9 disease relative to the susceptibility to said at least one physiological response, condition  
10 or disease in said standard.
- 11 19. The method of either of Claim 19 wherein said at least one addictive disease comprises  
12 opioid addiction; cocaine addiction or addiction to other psychostimulants; nicotine  
13 addiction; barbituate or sedative hypnotic addiction; anxiolytic addiction; or  
14 alcohol addiction.
- 15 20. A method for determining a susceptibility to pain in a subject relative to a susceptibility  
of pain in a standard, wherein the method comprises the steps of:  
a) removing a bodily sample from said subject, wherein said sample comprises a  
first and second allele comprising a human orphanin FQ/nociceptin receptor  
gene;  
b) determining whether said human orphanin FQ/nociceptin receptor gene of said  
first allele comprises a DNA sequence having at least one variation in SEQ ID  
NO:1, wherein said variation comprises G-46A, GIVS I 135C, GIVS I 250A,  
GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T, C1126G, or any  
combination thereof, such that the presence of said at least one variation in said  
human orphanin FQ/nociceptin receptor gene of said first allele is expected to be  
indicative of susceptibility to pain in said subject relative to susceptibility to  
pain in said standard, wherein said first allele of said standard comprises a  
human orphanin FQ/nociceptin receptor gene comprising a DNA sequence of  
SEQ ID NO:1.

- 1           21.    The method of Claim 20 for determining a susceptibility to pain in a subject, further  
2           comprising the step of determining whether said second allele of said bodily sample  
3           comprises a human orphanin FQ/nociceptin receptor gene comprising a DNA sequence  
4           having at least one variation in SEQ ID NO:1, wherein said variation comprises G-46A,  
5           GIVS I 135C, GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T,  
6           C1126G, or any combination thereof, such that the presence of said at least one variation  
7           in said second allele is expected to be indicative of susceptibility to pain in said subject  
8           relative to susceptibility of pain in said standard, wherein said second allele of said  
9           standard comprises a human orphanin FQ/nociceptin receptor gene comprising a DNA  
10          sequence of SEQ ID NO:1.
- 11          22.    A method for determining a therapeutically effective amount of pain reliever to  
            administer to a subject in order to induce analgesia in said subject relative to a  
            therapeutically effective amount of pain reliever to administer to a standard in order to  
            induce analgesia in said standard, wherein the method comprises determining a  
            susceptibility to pain in said subject relative to susceptibility to pain in said standard,  
            wherein susceptibility to pain in said subject is expected to be indicative of said  
            therapeutically effective amount of pain reliever to administer to said subject to induce  
            analgesia in said subject relative to said therapeutically effective amount of pain reliever  
            to administer to said standard to induce analgesia in said standard.
- 1          23.    The method of Claim 22 for determining a therapeutically effective amount of pain  
2           reliever to administer to said subject, wherein determining susceptibility to pain in said  
3           subject comprises the steps of:  
4           a)     removing a bodily sample from said subject, wherein said sample comprises a  
5                 first and second allele comprising a human orphanin FQ/nociceptin receptor  
6                 gene; and  
7           b)     determining whether said first allele comprises a human orphanin FQ/nociceptin  
8                 receptor gene comprising a DNA sequence having at least one variation in SEQ  
9                 ID NO:1, wherein said at least one variation comprises G-46A, GIVS I 135C,  
10                GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T, C1126G,  
11                or any combination thereof, wherein the presence of said at least one variation in

12 said human orphanin FQ/nociceptin receptor gene of said first allele is expected  
13 to be indicative of the subject's susceptibility to pain relative to said to  
14 susceptibility of pain in said standard, wherein said first allele of said standard  
15 comprises a human orphanin FQ/nociceptin receptor gene comprising a DNA  
16 sequence of SEQ ID NO:1, such that said therapeutically effective amount of  
17 pain reliever to administer to the subject in order to induce analgesia is related to  
18 said susceptibility to pain in said subject relative to susceptibility to pain in said  
19 standard.

1 24. The method of Claim 23, wherein determining susceptibility to pain in said subject  
2 relative to susceptibility to pain in said standard further comprises the step of  
3 determining whether said second allele of said bodily sample from said subject  
4 comprises a human orphanin FQ/nociceptin receptor gene comprising a DNA sequence  
5 having at least one variation in SEQ ID NO:1, wherein said at least one variation  
6 comprises G-46A, GIVS I 135C, GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T,  
7 A804G, C1026T, C1126G, or any combination thereof, such that the presence of said at  
8 least one variation in said second allele is expected to be indicative of susceptibility to  
9 pain in said subject relative to susceptibility to pain in said standard, wherein said second  
10 allele of said standard comprises a human orphanin FQ/nociceptin receptor gene  
11 comprising a DNA sequence of SEQ ID NO:1, and the therapeutically effective amount  
12 of pain reliever to administer to said subject to induce analgesia in said subject is related  
13 to the presence of said at least one variation in said human orphanin FQ/nociceptin  
14 receptor gene of said second allele of said bodily sample from said subject.

1 25. A method for determining a therapeutically effective amount of therapeutic agent to  
2 administer to a subject suffering from at least one addictive disease to treat the at least  
3 one addictive disease in said subject relative to a therapeutically effective amount of  
4 therapeutic agent to administer to a standard suffering from the at least one addictive  
5 disease to treat the at least one addictive disease in said standard, wherein the method  
6 comprises the steps of:  
7 a) removing a bodily sample from said subject, wherein said sample comprises a  
8 first and second allele comprising a human orphanin FQ/nociceptin receptor  
9 gene; and

b) determining whether said first allele comprises a human orphanin FQ/nociceptin receptor gene comprising a DNA sequence having at least one variation in SEQ ID NO:1, wherein said variation comprises G-46A, GIVS I 135C, GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T, C1126G, or any combination thereof,

wherein the presence of said at least one variation in said human orphanin FQ/nociceptin receptor gene of said first allele is expected to be indicative of the therapeutically effective amount of said therapeutic agent to administer to the subject to treat said at least one addictive disease in said subject relative to said therapeutically effective amount of said therapeutic agent to administer to said standard to treat said at least one addictive disease in said standard, wherein said first allele of said standard comprises a human orphanin FQ/nociceptin receptor gene comprising a DNA sequence of SEQ ID NO:1.

26. The method of Claim 25 for determining a therapeutically effective amount of therapeutic agent to administer to a subject suffering from said at least one addictive disease to treat said at least one addictive disease, relative to said therapeutically effective amount of said therapeutic agent administered to said standard suffering from said at least one addictive disease to treat said at least one addictive disease in said standard, further comprising the step of determining whether said second allele of said bodily sample from said subject comprises a human orphanin FQ/nociceptin receptor gene comprising a DNA sequence having at least one variation in SEQ ID NO:1, wherein said variation comprises G-46A, GIVS I 135C, GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T, C1126G, or any combination thereof, such that the presence of said at least one variation in said second allele related to said therapeutically effective amount of said therapeutic agent administered to said subject to treat said at least one addictive disease in said subject relative to said therapeutically effective amount of said therapeutic agent to administer to said standard to treat said at least one addictive disease in said standard, wherein said second allele of said standard comprises a human orphanin FQ/nociceptin receptor gene comprising a DNA sequence of SEQ ID NO:1.

27. The method of either of Claims 25 or 26, wherein said at least one addictive disease

comprises opioid addiction; cocaine addiction or addiction to other psychostimulants; nicotine addiction; barbiturate or sedative hypnotic addiction; anxiolytic addiction; or alcohol addiction.

28. A commercial test kit may for determining the presence of at least one variation in a human orphanin FQ/nociceptin receptor gene of an allele in a bodily sample taken from a subject, wherein the commercial test kit comprises:
  - a) PCR oligonucleotide primers suitable for detection of an allele comprising a human orphanin FQ/nociceptin receptor gene comprising a DNA sequence having at least one variation in SEQ ID NO:1 comprising G-46A, GIVS I 135C, GIVS I 250A, GIVS I 251A, C510T, CIVS III 67T, A804G, C1026T, C1126G, or any combination thereof;
  - b) other reagents; and
  - c) directions for use of the kit.
29. A nucleic acid comprising an intron of the human orphanin FQ/nociceptin receptor gene as set forth in SEQ ID No:2.
30. A nucleic acid as set forth in SEQ ID No:2.